

Int net

02/07/2006 10661109.trn

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NEWS 3 DEC 05 CASREACT(R) - Over 10 million reactions available
NEWS 4 DEC 14 2006 MeSH terms loaded in MEDLINE/LMEDLINE
NEWS 5 DEC 14 2006 MeSH terms loaded for MEDLINE file segment of TOXCENTER
NEWS 6 DEC 14 CA/CAPLUS to be enhanced with updated IPC codes
NEWS 7 DEC 21 IPC search and display fields enhanced in CA/CAPLUS with the
IPC reform
NEWS 8 DEC 23 New IPC8 SEARCH, DISPLAY, and SELECT fields in USPATFULL/
USPAT2
NEWS 9 JAN 13 IPC 8 searching in IFIPAT, IFIUDB, and IFICDB
NEWS 10 JAN 13 New IPC 8 SEARCH, DISPLAY, and SELECT enhancements added to
INPADOC
NEWS 11 JAN 17 Pre-1988 INPI data added to MARPAT
NEWS 12 JAN 17 IPC 8 in the WPI family of databases including WPIFV
NEWS 13 JAN 30 Saved answer limit increased
NEWS 14 JAN 31 Monthly current-awareness alert (SDI) frequency
added to TULSA

NEWS EXPRESS JANUARY 03 CURRENT VERSION FOR WINDOWS IS V8.01,
CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
AND CURRENT DISCOVER FILE IS DATED 19 DECEMBER 2005.
V8.0 USERS CAN OBTAIN THE UPGRADE TO V8.01 AT
<http://download.cas.org/express/v8.0-Discover/>

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NEWS PHONE Direct Dial and Telecommunication Network Access to STN
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FILE 'HOME' ENTERED AT 13:46:47 ON 07 FEB 2006

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COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

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STRUCTURE FILE UPDATES: 6 FEB 2006 HIGHEST RN 873652-66-5
DICTIONARY FILE UPDATES: 6 FEB 2006 HIGHEST RN 873652-66-5

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TSCA INFORMATION NOW CURRENT THROUGH JULY 14, 2005

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*
* The CA roles and document type information have been removed from *
* the IDE default display format and the ED field has been added, *
* effective March 20, 2005. A new display format, IDERL, is now *
* available and contains the CA role and document type information. *
*

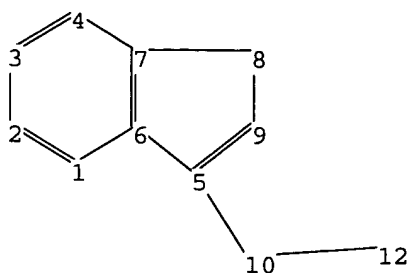
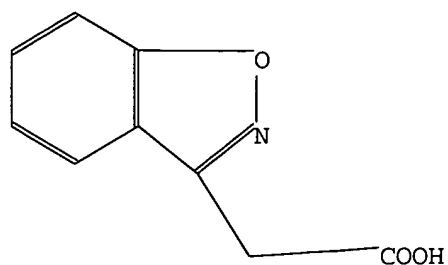
Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>

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```

chain nodes :
10 12
ring nodes :
1 2 3 4 5 6 7 8 9
chain bonds :
5-10 10-12
ring bonds :
1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9
exact/norm bonds :
5-9
exact bonds :
5-6 5-10 7-8 8-9 10-12
normalized bonds :
1-2 1-6 2-3 3-4 4-7 6-7
isolated ring systems :
containing 1 :

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Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
12:CLASS

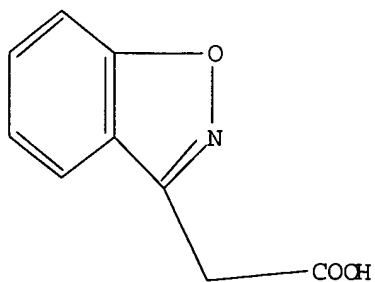
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L1 STRUCTURE UPLOADED

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=> D L1
L1 HAS NO ANSWERS
L1 STR

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Structure attributes must be viewed using STN Express query preparation.

=> S L1

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SAMPLE SEARCH INITIATED 13:47:16 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 23 TO ITERATE

100.0% PROCESSED 23 ITERATIONS 5 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 173 TO 747
PROJECTED ANSWERS: 5 TO 234

L2 5 SEA SSS SAM L1

=> S L1 SSS FULL
FULL SEARCH INITIATED 13:47:23 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 345 TO ITERATE

100.0% PROCESSED 345 ITERATIONS
SEARCH TIME: 00.00.01

115 ANSWERS

L3 115 SEA SSS FUL L1

=> S L3/P
'P' IS NOT A VALID CROSSOVER QUALIFIER FOR L3
Answer sets created in a different file may be field qualified with a
limited set of qualifiers. Enter HELP CROSSOVER at an arrow prompt
(=>) for specific information.

=> FIL HCAPLUS
COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE	TOTAL
ENTRY	SESSION
167.38	167.59

FILE 'HCAPLUS' ENTERED AT 13:47:59 ON 07 FEB 2006
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FILE COVERS 1907 - 7 Feb 2006 VOL 144 ISS 7
FILE LAST UPDATED: 6 Feb 2006 (20060206/ED)

New CAS Information Use Policies, enter HELP USAGETERMS for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L3
L4

62 L3

10661109.trn

Page 4

13:52

=> S L3/P

L5 26 L3/P

=> S L4 AND COUMARIN

23032 COUMARIN

6821 COUMARINS

24693 COUMARIN

(COUMARIN OR COUMARINS)

L6 5 L4 AND COUMARIN

=> S L5 AND COUMARIN

23032 COUMARIN

6821 COUMARINS

24693 COUMARIN

(COUMARIN OR COUMARINS)

L7 3 L5 AND COUMARIN

=> d l6 ibib abs hitstr tot

L6 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:429406 HCAPLUS

DOCUMENT NUMBER: 142:482033

TITLE: A process for the manufacture of zonisamide, useful as anticonvulsant agent

INVENTOR(S): Jaweed Mukarram, Siddiqui Mohammed; Merwade, Aravind Yehanathsa; Shukla, Jagdish Dattopant; Saiyad, Anis Mushtaqeali

PATENT ASSIGNEE(S): Wockhardt Limited, India

SOURCE: PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

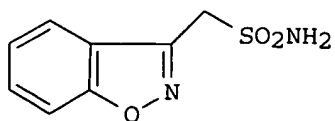
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005044808	A1	20050519	WO 2003-IB5052	20031111
W: AE, AG, AL, AM, <u>AT</u> , AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRIORITY APPLN. INFO.:			WO 2003-IB5052	20031111

OTHER SOURCE(S): CASREACT 142:482033

GI



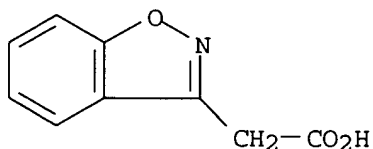
I

AB The invention relates to an improved process for the preparation of zonisamide (I), a well known anticonvulsant. Other aspects of this invention are isolation of a key intermediate, viz., isolation of crystalline sodium chloride associated with 1,2-benzisoxazole-3-methane sodium sulfonate (BOS-Na:NaCl). Zonisamide (I, 99% HPLC purity) was prepared via ring opening/cyclization of 4-hydroxycoumarin in the presence of NH₂OH (step 1), sulfonation of the obtained 1,2-benzisoxazole-3-acetic acid, and chlorination/amidation of the obtained sodium 1,2-benzisoxazole-3-methanesulfonate associated with NaCl (yield of step 1 was 95-98%). The anal. characteristics like IR and XRD data of BOS-Na:NaCl were also reported to confirm its nature.

IT **4865-84-3P**, 1,2-Benzisoxazole-3-acetic acid
 RL: IMF (Industrial manufacture); RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)
 (process for the manufacture of zonisamide useful as anticonvulsant agent)

RN 4865-84-3 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid (7CI, 8CI, 9CI) (CA INDEX NAME)



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:695963 HCAPLUS

DOCUMENT NUMBER: 137:216942

TITLE: Process for the preparation of 1,2-benzisoxazole-3-acetic acid, an intermediate in the synthesis of zonisamide

INVENTOR(S): Mendelovici, Mariofara, Nidam, Tamar

PATENT ASSIGNEE(S): Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals USA, Inc.

SOURCE: PCT Int. Appl., 14 pp.
 CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002070495	A1	20020912	WO 2002-US6419	20020304
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

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CA 2440030

AA 20020912

CA 2002-2440030

20020304

US 2002183525

A1 20021205

US 2002-90710

20020304

US 6677458

B2 20040113

EP 1373229

A1 20040102

EP 2002-717527

20020304

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

US 2004049053

A1 20040311

US 2003-661109

20030912

PRIORITY APPLN. INFO.:

US 2001-273172P

P 20010302

US 2001-294847P

P 20010531

US 2002-90710

A3 20020304

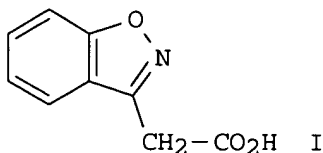
WO 2002-US6419

W 20020304

OTHER SOURCE(S):

CASREACT 137:216942

GI



AB A process for the preparation of 1,2-benzisoxazole-3-acetic acid (I) from 4-hydroxycoumarin and hydroxylamine.HCl in the presence of a base is disclosed. Compound I has com. importance as a key intermediate in the preparation of Zonisamide. For example, a solution of 4-hydroxycoumarin (100

g), hydroxylamine hydrochloride (150 g) and diethylamine (160 g) in MeOH (500 mL) was heated at reflux for 1 h. The reaction mixture was evaporated to dryness and the solid dissolved in aqueous NaHCO₃ and extracted with ether.

After

acidification of the aqueous phase, the product was isolated by filtration, washed with water and dried to provide I (99.82 g) in 93 % weight/weight yield. Advantages of the present invention are: (1) the prepare of I without the use of metallic sodium; and (2) the minimization of reaction side-products, e.g., oxime. The process is thus substantially less hazardous than previous methods. The invention also claims the prepare I or salts of which are converted to 1,2-benzisoxazole-3-methanesulfonamide, i.e., zonisamide.

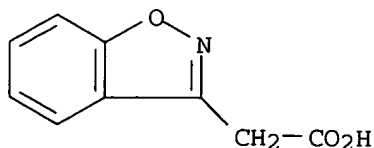
IT **4865-84-3P**, 1,2-Benzisoxazole-3-acetic acid

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(product; process for preparation of 1,2-benzisoxazole-3-acetic acid, an intermediate in synthesis of zonisamide)

RN 4865-84-3 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid (7CI, 8CI, 9CI) (CA INDEX NAME)



REFERENCE COUNT:

3

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

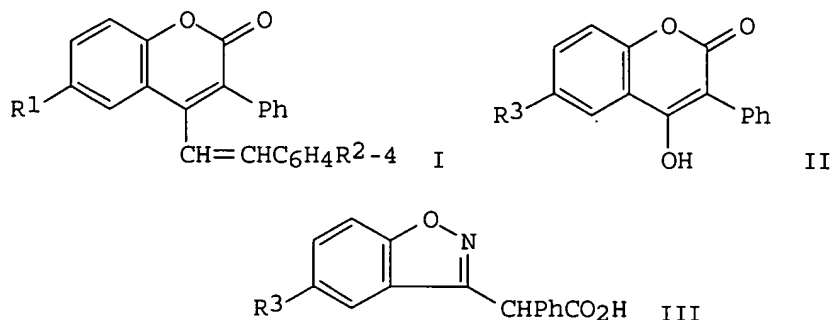
L6 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN

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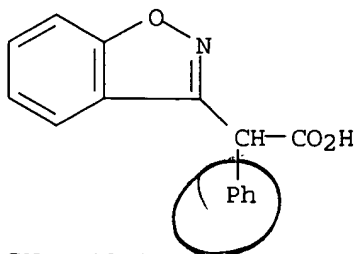
Page 7

13:52

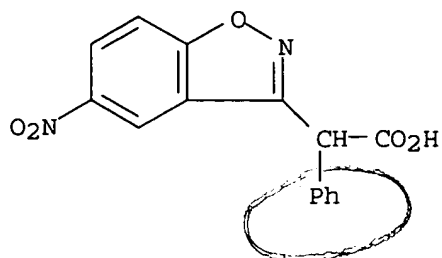
ACCESSION NUMBER: 1990:98332 HCAPLUS
 DOCUMENT NUMBER: 112:98332
 TITLE: Novel method in synthesis of 3-phenyl-4-styryl- and
 3-phenyl-4-hydroxycoumarins. Formation of
 3-phenylacetic acid benzisoxazole from
 3-phenyl-4-hydroxycoumarin and hydroxylamine
 hydrochloride
 AUTHOR(S): Lokhande, P. D.; Ghiya, B. J.
 CORPORATE SOURCE: Dep. Org. Chem., Inst. Sci., Nagpur, 440-001, India
 SOURCE: Journal of the Indian Chemical Society (1989), 66(5),
 314-15
 CODEN: JICSAH; ISSN: 0019-4522
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 OTHER SOURCE(S): CASREACT 112:98332
 GI



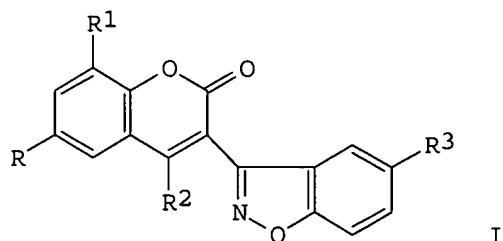
AB 2'-Hydroxychalcones were acylated by $\text{PhCH}_2\text{CO}_2\text{H}$, and the ester products were treated with KOH to give **coumarins** I ($\text{R}_1 = \text{H}, \text{Me}, \text{Cl}$; $\text{R}_2 = \text{OMe}, \text{H}$). Similarly, salicylate esters were converted to hydroxycoumarins II ($\text{R}_3 = \text{H}, \text{NO}_2$) which reacted with $\text{HONH}_2 \cdot \text{HCl}$ to give benzisoxazoles III.
 IT 125343-99-9 125344-00-5
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (ring contraction by, of hydroxycoumarins, benzisoxazoles from)
 RN 125343-99-9 HCAPLUS
 CN 1,2-Benzisoxazole-3-acetic acid, α -phenyl- (9CI) (CA INDEX NAME)



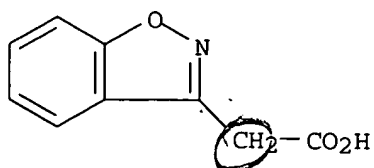
RN 125344-00-5 HCAPLUS
 CN 1,2-Benzisoxazole-3-acetic acid, 5-nitro- α -phenyl- (9CI) (CA INDEX NAME)



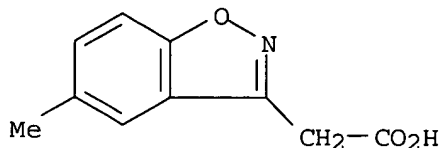
L6 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1979:203921 HCAPLUS
 DOCUMENT NUMBER: 90:203921
 TITLE: Synthesis of 3-(3'-benzisoxazolyl) coumarins
 AUTHOR(S): Lakshmi, A. Sree; Rao, K. Venkateswara; Sundaramurthy, V.
 CORPORATE SOURCE: Dep. Chem., Osmania Univ., Hyderabad, India
 SOURCE: Current Science (1979), 48(4), 153-4
 CODEN: CUSCAM; ISSN: 0011-3891
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI



AB ~~The cyclocondensation of benzisoxazole-3-acetic acids with salicylaldehydes and o-hydroxyacetophenones gave title compds. I (R = H, NO2, Cl, Me; R1 = H, NO2, Cl; R2 = H, Me; R3 = Me, H), useful as bactericides and fungicides (no data). Salicylaldehyde was treated with 5-methylbenzisoxazole-3-acetic acid, Ac2O, and Et3N to give I (R = R1 = R2 = H, R3 = Me).~~
 IT 4865-84-3 70154-01-7
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (cyclocondensation reaction with salicylaldehydes and 2'-hydroxyacetophenones)
 RN 4865-84-3 HCAPLUS
 CN 1,2-Benzisoxazole-3-acetic acid (7CI, 8CI, 9CI) (CA INDEX NAME)

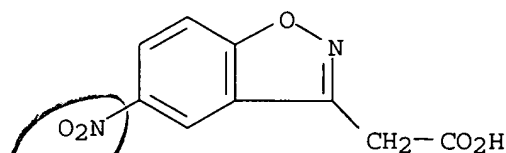


RN 70154-01-7 HCAPLUS
CN 1,2-Benzisoxazole-3-acetic acid, 5-methyl- (9CI) (CA INDEX NAME)

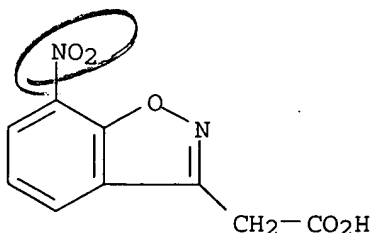


RSI. A518

L6 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1971:463554 HCAPLUS
DOCUMENT NUMBER: 75:63554
TITLE: Reaction between coumarins and hydroxylamine
AUTHOR(S): Giannella, Mario; Gualtieri, Fulvio; ~~Stein, Maria Luisa~~
CORPORATE SOURCE: Inst. Pharm. Org. Chem., Univ. Camerino, Camerino, Italy
SOURCE: Journal of Heterocyclic Chemistry (1971), 8(3), 397-403
CODEN: JHTCAD; ISSN: 0022-152X
DOCUMENT TYPE: Journal
LANGUAGE: English
AB The Posner reaction between coumarin and hydroxylamine was studied and extended to several substituted coumarins. Isolation of some significant reaction intermediates permitted rationalization of a possible reaction pathway.
IT 32906-16-4P 32906-17-5P 33026-03-8P 34610-57-6P
RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)
RN 32906-16-4 HCAPLUS
CN 1,2-Benzisoxazole-3-acetic acid, 5-nitro- (8CI, 9CI) (CA INDEX NAME)



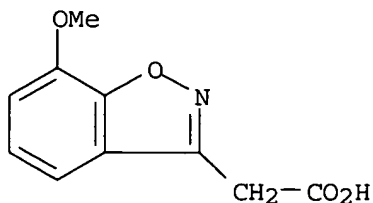
RN 32906-17-5 HCAPLUS
CN 1,2-Benzisoxazole-3-acetic acid, 7-nitro- (8CI, 9CI) (CA INDEX NAME)



RN 33026-03-8 HCAPLUS

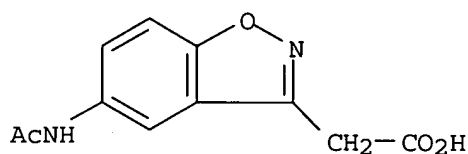
02/07/2006 10661109.trn

CN 1,2-Benzisoxazole-3-acetic acid, 7-methoxy- (8CI, 9CI) (CA INDEX NAME)



RN 34610-57-6 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid, 5-acetamido- (8CI) (CA INDEX NAME)



=> d 17 ibib abs hitstr tot

L7 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:429406 HCAPLUS

DOCUMENT NUMBER: 142:482033

TITLE: A process for the manufacture of zonisamide, useful as anticonvulsant agent

INVENTOR(S): Jaweed Mukarram, Siddiqui Mohammed; Merwade, Aravind Yehanathsa; Shukla, Jagdish Dattopant; Saiyad, Anis Mushtageali

PATENT ASSIGNEE(S): Wockhardt Limited, India

SOURCE: PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005044808	A1	20050519	WO 2003-IB5052	20031111
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RW:	BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

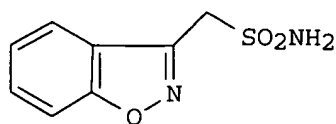
PRIORITY APPLN. INFO.:

WO 2003-IB5052

20031111

OTHER SOURCE(S): CASREACT 142:482033

GI



I

AB The invention relates to an improved process for the preparation of zonisamide (I), a well known anticonvulsant. Other aspects of this invention are isolation of a key intermediate, viz., isolation of crystalline sodium chloride associated with 1,2-benzisoxazole-3-methane sodium sulfonate (BOS-Na:NaCl). Zonisamide (I, 99% HPLC purity) was prepared via ring opening/cyclization of 4-hydroxycoumarin in the presence of NH_2OH (step 1), sulfonation of the obtained 1,2-benzisoxazole-3-acetic acid, and chlorination/amidation of the obtained sodium 1,2-benzisoxazole-3-methanesulfonate associated with NaCl (yield of step 1 was 95-98%). The anal. characteristics like IR and XRD data of BOS-Na:NaCl were also reported to confirm its nature.

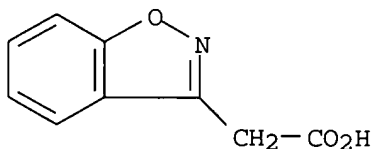
IT **4865-84-3P**, 1,2-Benzisoxazole-3-acetic acid

RL: IMF (Industrial manufacture); RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)

(process for the manufacture of zonisamide useful as anticonvulsant agent)

RN 4865-84-3 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid (7CI, 8CI, 9CI) (CA INDEX NAME)



REFERENCE COUNT:

1

THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:695963 HCAPLUS

DOCUMENT NUMBER: 137:216942

TITLE: Process for the preparation of 1,2-benzisoxazole-3-acetic acid, an intermediate in the synthesis of zonisamide

INVENTOR(S): Mendelovici, Mariorara; Nidam, Tamar

PATENT ASSIGNEE(S): Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals USA, Inc.

SOURCE: PCT Int. Appl., 14 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

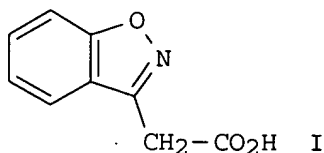
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002070495	A1	20020912	WO 2002-US6419	20020304
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,				

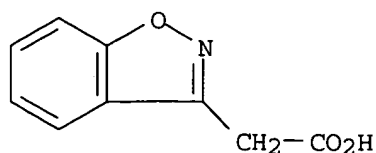
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
 PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
 UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
 TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
 CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
 BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
 CA 2440030 AA 20020912 CA 2002-2440030 20020304
 US 2002183525 A1 20021205 US 2002-90710 20020304
 US 6677458 B2 20040113
 EP 1373229 A1 20040102 EP 2002-717527 20020304
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
 US 2004049053 A1 20040311 US 2003-661109 20030912
 PRIORITY APPLN. INFO.: US 2001-273172P P 20010302
 US 2001-294847P P 20010531
 US 2002-90710 A3 20020304
 WO 2002-US6419 W 20020304
 OTHER SOURCE(S): CASREACT 137:216942
 GI



AB A process for the preparation of 1,2-benzisoxazole-3-acetic acid (I) from 4-hydroxycoumarin and hydroxylamine.HCl in the presence of a base is disclosed. Compound I has com. importance as a key intermediate in the preparation of Zonisamide. For example, a solution of 4-hydroxycoumarin (100 g), hydroxylamine hydrochloride (150 g) and diethylamine (160 g) in MeOH (500 mL) was heated at reflux for 1 h. The reaction mixture was evaporated to dryness and the solid dissolved in aqueous NaHCO₃ and extracted with ether. After acidification of the aqueous phase, the product was isolated by filtration, washed with water and dried to provide I (99.82 g) in 93 % weight/weight yield. Advantages of the present invention are: (1) the prepare of I without the use of metallic sodium; and (2) the minimization of reaction side-products, e.g., oxime. The process is thus substantially less hazardous than previous methods. The invention also claims the prepare I or salts of which are converted to 1,2-benzisoxazole-3-methanesulfonamide, i.e., zonisamide.

IT **4865-84-3P**, 1,2-Benzisoxazole-3-acetic acid
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
 (product; process for preparation of 1,2-benzisoxazole-3-acetic acid, an intermediate in synthesis of zonisamide)

RN 4865-84-3 HCAPLUS
 CN 1,2-Benzisoxazole-3-acetic acid (7CI, 8CI, 9CI) (CA INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1971:463554 HCAPLUS

DOCUMENT NUMBER: 75:63554

TITLE: Reaction between **coumarins** and hydroxylamine

AUTHOR(S): Giannella, Mario; Gualtieri, Fulvio; Stein, Maria Luisa

CORPORATE SOURCE: Inst. Pharm. Org. Chem., Univ. Camerino, Camerino, Italy

SOURCE: Journal of Heterocyclic Chemistry (1971), 8(3), 397-403

CODEN: JHTCAD; ISSN: 0022-152X

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The Posner reaction between **coumarin** and hydroxylamine was studied and extended to several substituted **coumarins**.

Isolation of some significant reaction intermediates permitted rationalization of a possible reaction pathway.

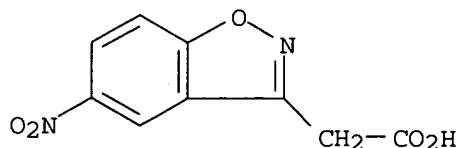
IT 32906-16-4P 32906-17-5P 33026-03-8P

34610-57-6P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)

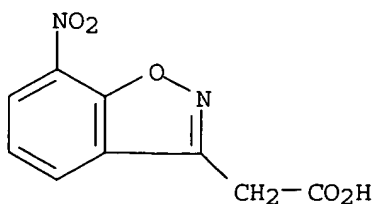
RN 32906-16-4 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid, 5-nitro- (8CI, 9CI) (CA INDEX NAME)



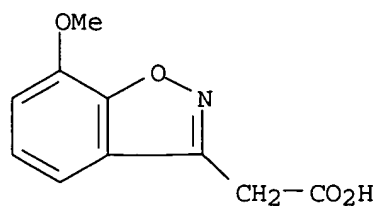
RN 32906-17-5 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid, 7-nitro- (8CI, 9CI) (CA INDEX NAME)

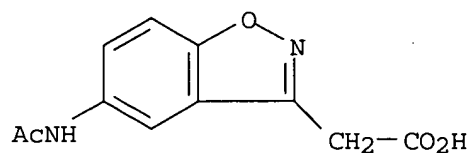


RN 33026-03-8 HCAPLUS

CN 1,2-Benzisoxazole-3-acetic acid, 7-methoxy- (8CI, 9CI) (CA INDEX NAME)



RN 34610-57-6 HCAPLUS
CN 1,2-Benzisoxazole-3-acetic acid, 5-acetamido- (8CI) (CA INDEX NAME)



=> S BENZISOXAZOLE-3-ACETIC ACID
1136 BENZISOXAZOLE
359 BENZISOXAZOLES
1217 BENZISOXAZOLE
(BENZISOXAZOLE OR BENZISOXAZOLES)
6487553 3
219947 ACETIC
22 ACETICS
219956 ACETIC
(ACETIC OR ACETICS)
4097027 ACID
1509527 ACIDS
4583575 ACID
(ACID OR ACIDS)
L8 38 BENZISOXAZOLE-3-ACETIC ACID
(BENZISOXAZOLE (W) 3 (W) ACETIC (W) ACID)

=> S L8 AND 4-HYDROXY-COUMARIN
5259858 4
434374 HYDROXY
9 HYDROXIES
434383 HYDROXY
(HYDROXY OR HYDROXIES)
23032 COUMARIN
6821 COUMARINS
24693 COUMARIN
(COUMARIN OR COUMARINS)
99 4-HYDROXY-COUMARIN
(4 (W) HYDROXY (W) COUMARIN)

L9 1 L8 AND 4-HYDROXY-COUMARIN

=> d-19 1b1b abs hitstr tot

L9 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2002:695963 HCAPLUS
DOCUMENT NUMBER: 137:216942

TITLE: Process for the preparation of 1,2-
benzisoxazole-3-acetic
acid, an intermediate in the synthesis of
 zonisamide

INVENTOR(S): Mendelovici, Mariorara; Nidam, Tamar

PATENT ASSIGNEE(S): Teva Pharmaceutical Industries Ltd., Israel; Teva
 Pharmaceuticals USA, Inc.

SOURCE: PCT Int. Appl., 14 pp.
 CODEN: PIXXD2

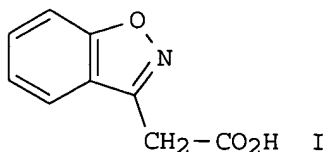
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002070495	A1	20020912	WO 2002-US6419	20020304
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2440030	AA	20020912	CA 2002-2440030	20020304
US 2002183525	A1	20021205	US 2002-90710	20020304
US 6677458	B2	20040113		
EP 1373229	A1	20040102	EP 2002-717527	20020304
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
US 2004049053	A1	20040311	US 2003-661109	20030912
PRIORITY APPLN. INFO.:				
			US 2001-273172P	P 20010302
			US 2001-294847P	P 20010531
			US 2002-90710	A3 20020304
			WO 2002-US6419	W 20020304
OTHER SOURCE(S): CASREACT 137:216942				
GI				



AB A process for the preparation of 1,2-**benzisoxazole-3-acetic acid** (I) from 4-hydroxycoumarin and hydroxylamine.HCl in the presence of a base is disclosed. Compound I has com. importance as a key intermediate in the preparation of Zonisamide. For example, a solution of 4-hydroxycoumarin (100 g), hydroxylamine hydrochloride (150 g) and diethylamine (160 g) in MeOH (500 mL) was heated at reflux for 1 h. The reaction mixture was evaporated to dryness and the solid dissolved in aqueous NaHCO₃ and extracted with ether. After acidification of the aqueous phase,

the product was isolated by filtration, washed with water and dried to provide I (99.82 g) in 93 % weight/weight yield. Advantages of the present invention are: (1) the prepare of I without the use of metallic sodium; and (2) the minimization of reaction side-products, e.g., oxime. The process is thus substantially less hazardous than previous methods. The invention also claims the prepare I or salts of which are converted to 1,2-benzisoxazole-3-methanesulfonamide, i.e., zonisamide.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> S L8 AND PROCESS
2199424 PROCESS
1480808 PROCESSES
3278489 PROCESS
(PROCESS OR PROCESSES)
L10 7 L8 AND PROCESS

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L10 ANSWER 1 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1050940 HCAPLUS

DOCUMENT NUMBER: 143:326350

TITLE: One-pot **process** for the preparation of 1,2-benzisoxazole-3-methanesulfonamide from 4-hydroxycoumarin

INVENTOR(S): Ueno, Yoshikazu; Ishikura, Tsutomu

PATENT ASSIGNEE(S): Japan

SOURCE: U.S. Pat. Appl. Publ., 5 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005215796	A1	20050929	US 2005-88802	20050325
WO 2005092869	A1	20051006	WO 2005-JP5349	20050324
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			

PRIORITY APPLN. INFO.: US 2004-556073P P 20040325

OTHER SOURCE(S): CASREACT 143:326350

AB 1,2-Benzisoxazole-3-methanesulfonamide was prepared by reaction of 4-hydroxycoumarin and NH₂OH (salt) in H₂O to give a mixture, acidification of the mixture and addition of ClCH₂CH₂Cl, removal of the aqueous layer to give a mixture containing 1,2-benzisoxazole-3-acetic acid and ClCH₂CH₂Cl, further removal of H₂O by distillation, addition of ClSO₃H, addition of base to give an alkali metal salt of 1,2-benzisoxazole-3-

methanesulfonic acid, addition of POCl₃ to give 1,2-benzisoxazole-3-methanesulfonyl chloride, and addition of NH₃.

L10 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:429406 HCAPLUS

DOCUMENT NUMBER: 142:482033

TITLE: A **process** for the manufacture of zonisamide, useful as anticonvulsant agent

INVENTOR(S): Jaweed Mukarram, Siddiqui Mohammed; Merwade, Aravind Yehanathsa; Shukla, Jagdish Dattopant; Saiyad, Anis Mushtaqali

PATENT ASSIGNEE(S): Wockhardt Limited, India

SOURCE: PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

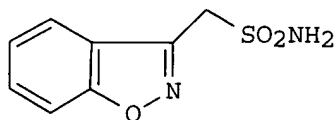
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005044808	A1	20050519	WO 2003-IB5052	20031111
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
PRIORITY APPLN. INFO.:			WO 2003-IB5052	20031111
OTHER SOURCE(S):	CASREACT 142:482033			

GI



AB The invention relates to an improved **process** for the preparation of zonisamide (I), a well known anticonvulsant. Other aspects of this invention are isolation of a key intermediate, viz., isolation of crystalline sodium chloride associated with 1,2-benzisoxazole-3-methane sodium sulfonate (BOS-Na:NaCl). Zonisamide (I, 99% HPLC purity) was prepared via ring opening/cyclization of 4-hydroxycoumarin in the presence of NH₂OH (step 1), sulfonation of the obtained 1,2-benzisoxazole-3-acetic acid, and chlorination/amidation of the obtained sodium 1,2-benzisoxazole-3-methanesulfonate associated with NaCl (yield of step 1 was 95-98%). The anal. characteristics like IR and XRD data of BOS-Na:NaCl were also reported to confirm its nature.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L10 ANSWER 3 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:590879 HCAPLUS
 DOCUMENT NUMBER: 139:154994
 TITLE: Novel sulfonation method for zonisamide intermediate in zonisamide synthesis and their novel crystal forms
 INVENTOR(S): Nidam, Tamar; Mendelovici, Marioara; Schwartz, Edward; Wizel, Shlomit
 PATENT ASSIGNEE(S): Israel
 SOURCE: U.S. Pat. Appl. Publ., 35 pp., Cont.-in-part of U.S. Ser. No. 233,190.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2003144527	A1	20030731	US 2002-288135	20021105
US 2003114682	A1	20030619	US 2002-233190	20020829
US 6841683	B2	20050111		
WO 2004020419	A1	20040311	WO 2002-US35537	20021105
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
US 2004138471	A1	20040715	US 2003-662966	20030915
US 2004138472	A1	20040715	US 2003-662986	20030915
US 2005027126	A1	20050203	US 2004-928313	20040830
PRIORITY APPLN. INFO.:			US 2001-316109P	P 20010830
			US 2001-344439P	P 20011024
			US 2002-233190	A2 20020829

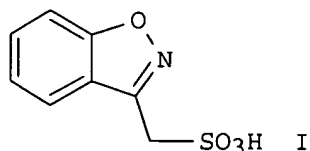
AB The present invention relates to a novel sulfonation of an intermediate of zonisamide. The sulfonation **processes** using chlorosulfonic acid as well as acetic anhydride and sulfuric acid in an organic solvent are disclosed. Crystalline forms of benzisoxazole methanesulfonic acid (BOS-H) and its salts (BOS-Na, BOS-Ca, and BOS-Ba) and their novel preparation **processes** are disclosed.

L10 ANSWER 4 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:202630 HCAPLUS
 DOCUMENT NUMBER: 138:221579
 TITLE: **Process** for the preparation of 1,2-benzisoxazole-3-methanesulfonic acid and its salts, intermediates in the synthesis of Zonisamide
 INVENTOR(S): Nidam, Tamar; Mendelovici, Marioara; Schwartz, Eduard; Wizel, Shlomit
 PATENT ASSIGNEE(S): Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals USA, Inc.
 SOURCE: PCT Int. Appl., 62 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent

LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003020708	A1	20030313	WO 2002-US27593	20020829
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2458905	AA	20030313	CA 2002-2458905	20020829
EP 1430037	A1	20040623	EP 2002-768748	20020829
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK				
JP 2005506980	T2	20050310	JP 2003-524979	20020829
PRIORITY APPLN. INFO.:			US 2001-316109P	P 20010830
			US 2001-344439P	P 20011024
			WO 2002-US27593	W 20020829
OTHER SOURCE(S):		CASREACT 138:221579		
GI				



AB A **process** for the preparation of 1,2-benzisoxazole-3-methanesulfonic acid (I) by sulfonation of 1,2-benzisoxazole-3-acetic acid with chlorosulfonic acid or acyl sulfates in an organic solvent and optional conversion to its salts is disclosed. I has com. importance as a key intermediate in the preparation of Zonisamide. For example, a solution of 1,2-benzisoxazole-3-acetic acid (20 gm), 98% H₂SO₄ (22 gm), and Ac₂O (23 gm) in AcOEt (80 mL) was heated at reflux for 4 h and the cooled reaction mixture treated with aqueous 10% aqueous NaOH (120 mL) to give I•Na (20.33 gm) in 100% purity. Advantages of the present invention are: (1) the preparation of I without the use of dioxane, improving the environmental safety of the reaction; and (2) the increased selectivity for preparation of the monosulfonated over the bisulfonated benzisoxazole. Crystalline forms of 1,2-benzisoxazole-3-methanesulfonic acid (BOS-H) and its salts (BOS-Na, BOS-Ca, and BOS-Ba) were also characterized.

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L10 ANSWER 5 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2002:695963 HCAPLUS
 DOCUMENT NUMBER: 137:216942

TITLE: **Process for the preparation of 1,2-benzisoxazole-3-acetic acid**, an intermediate in the synthesis of zonisamide

INVENTOR(S): Mendelovici, Mariorara; Nidam, Tamar

PATENT ASSIGNEE(S): Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals USA, Inc.

SOURCE: PCT Int. Appl., 14 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

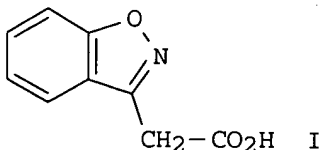
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002070495	A1	20020912	WO 2002-US6419	20020304
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
CA 2440030	AA	20020912	CA 2002-2440030	20020304
US 2002183525	A1	20021205	US 2002-90710	20020304
US 6677458	B2	20040113		
EP 1373229	A1	20040102	EP 2002-717527	20020304
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR			
US 2004049053	A1	20040311	US 2003-661109	20030912
PRIORITY APPLN. INFO.:			US 2001-273172P	P 20010302
			US 2001-294847P	P 20010531
			US 2002-90710	A3 20020304
			WO 2002-US6419	W 20020304

OTHER SOURCE(S): CASREACT 137:216942

GI



AB A **process** for the preparation of 1,2-benzisoxazole-3-acetic acid (I) from 4-hydroxycoumarin and hydroxylamine.HCl in the presence of a base is disclosed. Compound I has com. importance as a key intermediate in the preparation of Zonisamide. For example, a solution of 4-hydroxycoumarin (100 g), hydroxylamine hydrochloride (150 g) and diethylamine (160 g) in MeOH (500 mL) was heated at reflux for 1 h. The reaction mixture was evaporated to dryness and the solid dissolved in aqueous NaHCO₃ and extracted with ether. After acidification of the aqueous phase,

the product was isolated by filtration, washed with water and dried to provide I (99.82 g) in 93 % weight/weight yield. Advantages of the present invention are: (1) the prepare of I without the use of metallic sodium; and (2) the minimization of reaction side-products, e.g., oxime. The **process** is thus substantially less hazardous than previous methods. The invention also claims the prepare I or salts of which are converted to 1,2-benzisoxazole-3-methanesulfonamide, i.e., zonisamide.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L10 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1996:328908 HCAPLUS

DOCUMENT NUMBER: 125:53737

TITLE: Methoxylation modifies the activity of 1,2-**benzisoxazole-3-acetic acid**: 6,7-dimethoxy-1,2-**benzisoxazole-3-acetic acid** is an auxin antagonist in cytokinin mediated **processes**

AUTHOR(S): Ricci, Ada; Maggiali, Cesare Augusto; Torelli, Anna; Amorosi, Sonia; Ronchini, Ferdinando; Branca, Camillo
CORPORATE SOURCE: Dipartimento di Biologia Evolutiva, Via delle Scienze, Parma, 43100, Italy

SOURCE: Plant Science (Shannon, Ireland) (1996), 117(1,2), 151-158

CODEN: PLSCE4; ISSN: 0168-9452

PUBLISHER: Elsevier

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The insertion of a methoxy group in different positions of the aromatic ring modifies the activity of 1,2-**benzisoxazole-3-acetic acid** (BOAA), a specific morphogenetic compound with no activity on cell elongation or root growth. Monomethoxylation in the 4- and 7-position is critical in determining the kind of activity: 4-OMeBOAA induces stem elongation, inhibits root growth and does not improve shoot production; 7-OMeBOAA inhibits stem elongation and shoot production and is unable to induce root growth. 6,7-OMeBOAA, inactive on stem elongation and root growth, is unable to induce the expression of Pg5-GUS gene in the presence of BAP and inhibits the expression of this gene when induced by BAP plus IAA. Furthermore, 6,7-OMeBOAA inhibits completely shoot production and can therefore be regarded as an auxin antagonist in these cytokinin-mediated **processes**.

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ACCESSION NUMBER: 1990:547166 HCAPLUS

DOCUMENT NUMBER: 113:147166

TITLE: Effects of benzisoxazole and benzisothiazole on tomato plant regeneration in vitro

AUTHOR(S): Branca, Camillo; Torelli, Anna; Bassi, Maria

CORPORATE SOURCE: Ist. Bot., Univ. Parma, Parma, 43100, Italy

SOURCE: Plant Cell, Tissue and Organ Culture (1990), 21(1), 17-19

CODEN: PTCEDJ; ISSN: 0167-6857

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The effects of two synthetic auxins, 1,2-benzisothiazole-3-acetic acid (BOA) and 1,2-**benzisoxazole-3-acetic acid** (BIA), on plant regeneration in vitro have been studied on

explants of tomato cotyledons. The activity of these substances on cell elongation has also been tested on pea stem segments. BOA is particularly effective in inducing the formation of shoots but has a weak activity on cell elongation, while BIA, which is more effective in inducing cell elongation, is less active in morphogenesis. Thus, these two activities are not related to each other, the receptors involved in the two **processes** are probably different, and the chemical structure of the auxin may be an important factor in its morphogenetic action.

=> LOG Y

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
83.04	250.63

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
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